

L Number	Hits	Search Text	DB	Time stamp
1	1276	multiplex\$4 NEAR (pcr amplification)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:02
2	162	((multiplex\$4 NEAR (pcr amplification)) and (Invader or taqman))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 16:56
3	15	((multiplex\$4 NEAR (pcr amplification)) and (Invader or taqman)) and ((hot near start) or (taqstart))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 16:57
4	4	((multiplex\$4 NEAR (pcr amplification)) and (Invader or taqman)) and ((hot near start) or (taqstart)) and (SNP or (single adj1 nucleotide adj1 polymorphism))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 16:58
5	0	(multiplex\$4 NEAR (pcr amplification)) and (at adj1 least adj1 ("50" or fifty))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:03
6	35	(multiplex\$4 NEAR (pcr amplification)) and (at adj1 least adj5 primer)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:03
7	17	((multiplex\$4 NEAR (pcr amplification)) and (at adj1 least adj5 primer)) and (SNP or (single adj1 nucleotide adj1 polymorphism))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:06
8	18	((multiplex\$4 NEAR (pcr amplification)) and (at adj1 least adj5 primer)) not ((multiplex\$4 NEAR (pcr amplification)) and (at adj1 least adj5 primer)) and (SNP or (single adj1 nucleotide adj1 polymorphism))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:39
13	59		USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:37
14	14		USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:39
15	2	("6489455").PN.	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:39
16	0	((("6489455").PN.) and (SNP or (single adj1 nucleotide adj1 polymorphism)))	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/10/02 17:39

FILE 'BIOSIS, MEDLINE, EMBAL, EMBASE, SCISEARCH, BIOTECHDS, CAPLUS'
ENTERED AT 15:44:55 ON 02 OCT 2003

L1 40107 S SNP? OR (SINGLE (1W) NUCLEOTIDE (1W) POLYMORPHISM?)

L2 366 S L1 AND (INVADER? OR TAQMAN?)

L3 41 S L2 AND (MULTIPLEX)

L4 42 S L2 AND (MULTIPLEX?)

L5 0 S L4 AND (MULTIPLEX? (P) (AMPLIFICATION? PCR?))

L6 18 DUP REM L4 (24 DUPLICATES REMOVED)

L6 ANSWER 1 OF 18 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:248267 CAPLUS

DOCUMENT NUMBER: 138:379797

TITLE: Genome-wide association study of bronchial asthma
based on SNP analysis

AUTHOR(S): Tamati, Mayumi

CORPORATE SOURCE: Laboratory for Functional Analysis, SNP Research
Center, The Institute of Physical and Chemical
Research (RIKEN), Japan

SOURCE: Saishin Igaku (2003), 58(2), 209-215

CODEN: SAIGAK; ISSN: 0370-8241

PUBLISHER: Saishin Igakusha

DOCUMENT TYPE: Journal; General Review

LANGUAGE: Japanese

TI Genome-wide association study of bronchial asthma based on SNP
analysis

AB A review gives an overview of genome-wide genetic linkage study of
bronchial asthma based on SNP anal. Development of
establishment of database genome-wide SNP information for
Japanese population was summarized. Current status of the SNP
anal. and linkage anal. for identifying asthma-related genes in Japan was
discussed. The advantage of the use of the Multiplex PCR and
the Invader methods in collecting SNP information was
described.

ST review genome wide SNP genetic linkage analysis bronchial asthma

IT Databases

(SNP database; genome-wide assocn. study of bronchial asthma
based on SNP anal.)

IT Gene, animal

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(assocd. with bronchial asthma; genome-wide assocn. study of bronchial
asthma based on SNP anal.)

IT Asthma

Genetic linkage

Human

(genome-wide assocn. study of bronchial asthma based on SNP
anal.)

IT Genetic polymorphism

(single nucleotide; genome-wide assocn. study of bronchial asthma based on SNP anal.)

L6 ANSWER 2 OF 18 SCISEARCH COPYRIGHT 2003 THOMSON ISI on STN

ACCESSION NUMBER: 2003:70704 SCISEARCH

THE GENUINE ARTICLE: 634QF

TITLE: Improved sensitivity for solid-support invasive cleavage reactions with flow cytometry analysis

AUTHOR: Stevens P W (Reprint); Rao K V N; Hall J G; Lyamichev V; Neri B P; Kelso D M

CORPORATE SOURCE: Northwestern Univ, Robert R McCormick Sch Engn & Appl Sci,

Dept Biomed Engn, 2145 Sheridan Rd, Evanston, IL 60208 USA
(Reprint); Northwestern Univ, Robert R McCormick Sch Engn & Appl Sci, Dept Biomed Engn, Evanston, IL 60208 USA;
Third Wave Technol, Madison, WI USA

COUNTRY OF AUTHOR: USA

SOURCE: BIOTECHNIQUES, (JAN 2003) Vol. 34, No. 1, pp. 198-203.

Publisher: EATON PUBLISHING CO, 154 E. CENTRAL ST, NATICK, MA 01760 USA.

ISSN: 0736-6205.

DOCUMENT TYPE: Article; Journal

LANGUAGE: English

REFERENCE COUNT: 15

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB . . . nucleotide differences at target concentrations less than 200 fm. This sensitivity level is within the range required for analysis of SNPs in genomic DNA. In addition, the flow cytometry format has multiplexing potential, making the microsphere-based invasive cleavage assay attractive for high throughput genomic applications.

STP KeyWords Plus (R): SIGNAL AMPLIFICATION REACTION; OLIGONUCLEOTIDE PROBES;

INVADER ASSAY; IDENTIFICATION; DNA; PLATFORM

L6 ANSWER 3 OF 18 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:136073 CAPLUS

DOCUMENT NUMBER: 136:196564

TITLE: Non-fluorescent asymmetric cyanine dye compounds useful for quenching reporter dyes

INVENTOR(S): Lee, Linda G.; Graham, Ronald J.; Mullah, Khairuzzaman B.; Haxo, Francis T.

PATENT ASSIGNEE(S): PE Corporation (NY), USA

SOURCE: U.S., 62 pp., Cont.-in-part of U.S. 6,080,868.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6348596	B1	20020219	US 1999-357740	19990720
US 6080868	A	20000627	US 1998-12525	19980123
US 6541618	B1	20030401	US 2000-602544	20000621

PRIORITY APPLN. INFO.: US 1998-12525 A2 19980123
OTHER SOURCE(S): MARPAT 136:196564
REFERENCE COUNT: 84 THERE ARE 84 CITED REFERENCES
AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

AB The invention provides asym. cyanine dye compds. I, including substituted forms thereof, which are non-fluorescent quencher mols. The invention further provides reporter-quencher dye pairs, wherein the asym. cyanine dyes are the quenchers, polynucleotides incorporating the asym. cyanine dyes, and nucleic acid hybridization detection methods utilizing the dye-labeled polynucleotides. Nitrothiazole blue XXXIV was prep'd. from 2-methylbenzothiazole and used as a quencher dye paired with FAM or TET reporter dyes in Taqman assays.

ST asym cyanine dye fluorescence quenching reagent; nucleic acid hybridization asym cyanine dye fluorescence quencher; nitrothiazole blue quencher dye Taqman assay

IT PCR (polymerase chain reaction)
(Taqman assay, doubly-labeled probe for; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)

IT Gene, animal
RL: ANT (Analyte); ANST (Analytical study)
(for .beta.-actin of human, Taqman assay for; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)

IT PCR (polymerase chain reaction)
(multiplex; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)

IT Actins
RL: ANT (Analyte); ANST (Analytical study)
(.beta.-, Taqman assay for human gene for; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)

IT 400185-16-2 400185-17-3 400185-18-4 400185-19-5 400185-20-8
400185-21-9

RL: ARG (Analytical reagent use); PRP (Properties); ANST (Analytical study); USES (Uses)
(nucleotide sequence, SNP primer; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)

ACCESSION NUMBER: 2002:784266 CAPLUS
 DOCUMENT NUMBER: 137:305690
 TITLE: High-throughput single-nucleotide
 polymorphism typing system by
 multiplex PCR-Invader assay
 INVENTOR(S): Nakamura, Yusuke; Tanaka, Toshihiro; Onishi, Yozo;
 Ozaki, Koichi; Yamada, Akira
 PATENT ASSIGNEE(S): Institute of Physical and Chemical Research, Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 45 pp.
 CODEN: JKXXAF
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2002300894	A2	20021015	JP 2002-19752	20020129
US 2002182622	A1	20021205	US 2002-60301	20020201
PRIORITY APPLN. INFO.:			JP 2001-25700	A 20010201
TI High-throughput single-nucleotide polymorphism typing system by multiplex PCR-Invader assay				
AB A method for genotyping of single-nucleotide polymorphisms (SNPs) by nucleic acid amplification with a large no. of primers, is disclosed. Hot-start PCR, Taqman PCR, or invader method, may be preferably used. One of the most difficult issues to be solved in genome-wide assocn. studies is to reduce the amt. of genomic DNA required for genotyping. Currently available technologies require too large a quantity of genomic DNA to genotype with hundreds or thousands of single-nucleotide polymorphisms (SNPs). To overcome this problem, the authors combined the Invader assay with multiplex polymerase chain reaction (PCR), carried out in the presence of antibody to Taq polymerase, as well as using a novel 384-well card system that can reduce the required reaction vol. The authors amplified 100 genomic DNA fragments, each contg. one SNP, in a single tube, and analyzed each SNP with the Invader assay. This procedure correctly genotyped 98 of the 100 SNP loci examd. in PCR-amplified samples from ten individuals; the genotypes were confirmed by direct sequencing. The reproducibility and universality of the method were confirmed with two addnl. sets of 100 SNPs. Because the authors used 40ng of genomic DNA as a template for multiplex PCR, the amt. needed to assay one SNP was only 0.4 ng; therefore, theor., more than 200,000 SNPs could be genotyped at once when 100 .mu.g of genomic DNA is available. Our results indicate the feasibility of undertaking genome-wide assocn. studies using blood samples				

of only 5-10mL.

L6 ANSWER 5 OF 18 MEDLINE on STN DUPLICATE 1
ACCESSION NUMBER: 2002704237 MEDLINE
DOCUMENT NUMBER: 22353531 PubMed ID: 12465410
TITLE: A high-throughput SNP typing system for
 genome-wide association studies.
AUTHOR: Ohnishi Yozo
CORPORATE SOURCE: Laboratory for Cardiovascular Diseases, SNP Research
 Center, Institute of Physical and Chemical Research
 (RIKEN), 4-6-1 Shirokanedai, Minato-ku, Tokyo 108-8639,
 Japan.
SOURCE: GAN TO KAGAKU RYOHO [JAPANESE JOURNAL OF CANCER
AND
 CHEMOTHERAPY], (2002 Nov) 29 (11) 2031-6.
 Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200212
ENTRY DATE: Entered STN: 20021217
 Last Updated on STN: 20021218
 Entered Medline: 20021213
TI A high-throughput SNP typing system for genome-wide association
studies.
AB SNPs are useful markers for identifying genes responsible for
and/or associated with common diseases, and for directing personalized
medical care. Furthermore, because they are so frequent in the genome and
can be genotyped quite easily, SNPs can serve as markers for a
whole genome association study. However, one of the most difficult issues
to be solved for whole-genome association studies using SNPs is
reduction of the amount of genomic DNA for genotyping. The presently
available technologies require too much genomic DNA to be practical. To
overcome this problem, we combined the Invader assay with
multiplex PCR performed in the presence of Taq polymerase antibody
as well as a novel 384-well card system that reduces the reaction volume.
We amplified 96 genomic DNA fragments simultaneously in a single tube, and
analyzed each SNP using the Invader assay. Since we
used 10-20 nanograms of genomic DNA as a template for multiplex
PCR, the amount needed to assay one SNP was only 0.1-0.2
nanograms. Our results strongly indicate the feasibility of undertaking
genome-wide association studies using blood samples of only. . .

L6 ANSWER 6 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

ACCESSION NUMBER: 2002:616684 BIOSIS

DOCUMENT NUMBER: PREV200200616684

TITLE: A high-throughput SNP typing system for
genome-wide association studies in patients with myocardial
infarction.

AUTHOR(S): Ohnishi, Y. (1); Tanaka, T. (1); Ozaki, K. (1); Sekine, A.
(1); Nakamura, Y. (1)

CORPORATE SOURCE: (1) SNP Research Ctr, RIKEN, Tokyo Japan

SOURCE: American Journal of Human Genetics, (October, 2002) Vol.
71, No. 4 Supplement, pp. 435.

<http://www.journals.uchicago.edu/AJHG/home.html>. print.

Meeting Info.: 52nd Annual Meeting of the American Society
of Human Genetics Baltimore, MD, USA October 15-19, 2002
American Society of Human Genetics

. ISSN: 0002-9297.

DOCUMENT TYPE: Conference

LANGUAGE: English

TI A high-throughput SNP typing system for genome-wide association
studies in patients with myocardial infarction.

IT Methods & Equipment

Invader assay; assessment method; genotyping:

characterization method; high-throughput SNP typing system:

laboratory equipment; multiplex PCR

IT Miscellaneous Descriptors

genome-wide association; Meeting Abstract

L6 ANSWER 7 OF 18 BIOTECHDS COPYRIGHT 2003 THOMSON

DERWENT/ISI on STN

ACCESSION NUMBER: 2003-01265 BIOTECHDS

TITLE: Rapid multiplex single nucleotide
polymorphism genotyping based on single base
extension reactions and color-coded beads;
SNP genotyping using microphore bead washing
after polymerase chain reaction

AUTHOR: FUJIMURA N; KOHARA Y; OKANO K; YOHDA M; KAMBARA H

CORPORATE SOURCE: Tokyo Univ Agr and Technol; Hitachi Ltd

LOCATION: Kohara Y, Tokyo Univ Agr and Technol, Dept Biotechnol and
Life Sci, 2-24-16 Nakacho, Koganei, Tokyo 1840012, Japan

SOURCE: JOURNAL OF BIOSCIENCE AND BIOENGINEERING; (2002) 94,
4,

368-370

ISSN: 1389-1723

DOCUMENT TYPE: Journal

LANGUAGE: English

TI Rapid multiplex single nucleotide

polymorphism genotyping based on single base extension reactions

and color-coded beads;

SNP genotyping using microphore bead washing after
polymerase chain reaction

AB AUTHOR ABSTRACT - A Single nucleotide
polymorphism (SNP) typing method using color-coded
beads is promising because it is easy to use and inexpensive. However,
the present protocols are. . . diagnostic applications because they
need centrifugation for bead-washing. Here, we developed a simplified
protocol without a bead-washing procedure that enables SNP
typing of PCR amplified fragments in only 30 min.

DERWENT ABSTRACT: SNPs are the most frequent DNA polymorphism
found. They are expected to be diagnosis markers for a number of diseases
and drug responses. Many SNP typing methods, including the
TaqMan assay, Invader assay, molecular beacon,
polymerase chain reaction and electrophoresis, mass spectrometry,
pyro-DNA sequencing, bioluminometric assay coupled with modified primer
extension reactions, DNA chip, and bead technology, have been developed.
The paper looks at a rapid and simplified SNP typing method
using single base extension reactions and color-coded beads(3 pages)

CT SNP, GENOTYPING, SINGLE BASE EXTENSION, POLYMERASE CHAIN
REACTION, MICROSPHERE BEAD, DNA PRIMER, FLOW CYTOMETRY
ANALYSIS, APPL.

DIAGNOSIS DNA AMPLIFICATION HYBRIDIZATION (22,. . .

L6 ANSWER 8 OF 18 EMBASE COPYRIGHT 2003 ELSEVIER INC. ALL RIGHTS
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on STN

ACCESSION NUMBER: 2002085075 EMBASE

TITLE: High-throughput SNP genotyping.

AUTHOR: Jenkins S.; Gibson N.

CORPORATE SOURCE: S. Jenkins, R and D Genetics, AstraZeneca, Mereside,
Alderley Park, Macclesfield, Cheshire SK10 4TG, United
Kingdom. suzanne.jenkins@astrazeneca.com

SOURCE: Comparative and Functional Genomics, (2002) 3/1 (57-66).
Refs: 46

ISSN: 1531-6912 CODEN: YESTE3

COUNTRY: United Kingdom

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 022 Human Genetics

027 Biophysics, Bioengineering and Medical
Instrumentation

LANGUAGE: English

SUMMARY LANGUAGE: English

TI High-throughput SNP genotyping.

AB Whole genome approaches using single nucleotide
polymorphism (SNP) markers have the potential to

transform complex disease genetics and expedite pharmacogenetics research. This has led to a requirement for high-throughput SNP genotyping platforms. Development of a successful high-throughput genotyping platform depends on coupling reliable assay chemistry with an appropriate detection system. . . only a few cents per genotype. In addition, DNA template requirements must be minimised such that hundreds of thousands of SNPs can be interrogated using a relatively small amount of genomic DNA. As such, it is predicted that the next generation of high-throughput genotyping platforms will exploit large-scale multiplex reactions and solid phase assay detection systems.
Copyright .COPYRG.T. 2001 John Wiley & Sons, Ltd.

L6 ANSWER 9 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2003:390975 BIOSIS

DOCUMENT NUMBER: PREV200300390975

TITLE: A multiplex solution for SNP
interrogation using fifth-dye labeled short size standard
and single-tube primer extension reaction.

AUTHOR(S): Kuo, S. S. (1); Wei, D. (1); Ayanoglu, G. (1); Johnson, M.
(1); Chang, C. (1); Hanachi, P. (1); Tomaney, A. B. (1);
Dong, P. (1)

CORPORATE SOURCE: (1) Applied Biosystems, Foster City, CA, USA:
kuoss@appliedbiosystems.com USA

SOURCE: European Journal of Human Genetics, (2001) Vol. 9, No.
Supplement 1, pp. P1238. print.
Meeting Info.: 10th International Congress of Human
Genetics Vienna, Austria May 15-19, 2001 International
Federation of Human Genetics Societies
. ISSN: 1018-4813.

DOCUMENT TYPE: Conference

LANGUAGE: English

TI A multiplex solution for SNP interrogation using
fifth-dye labeled short size standard and single-tube primer extension
reaction.

IT . . .
techniques, laboratory techniques; fifth-dye labeled short size
standard extension reaction: genetic techniques, laboratory techniques;
genotyping: genetic techniques, laboratory techniques; real-time
Taqman PCR analysis [real-time Taqman polymerase
chain reaction analysis]: genetic techniques, laboratory techniques;
single-tube primer extension reaction: laboratory techniques

IT Miscellaneous Descriptors

SNP [single nucleotide
polymorphisms]; multiplex solution; Meeting Abstract

L6 ANSWER 10 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

DUPLICATE 2

ACCESSION NUMBER: 2001:409593 BIOSIS

DOCUMENT NUMBER: PREV200100409593

TITLE: A high-throughput SNP typing system for
genome-wide association studies.

AUTHOR(S): Ohnishi, Yozo; Tanaka, Toshihiro; Ozaki, Kouichi; Yamada,
Ryo; Suzuki, Hideyuki; Nakamura, Yusuke (1)

CORPORATE SOURCE: (1) Laboratory of Molecular Medicine, Human Genome
Center,

Institute of Medical Science, University of Tokyo, 4-6-1
Shirokanedai, Minato-ku, Tokyo, 108-8639:
yusuke@ims.u-tokyo.ac.jp Japan

SOURCE: Journal of Human Genetics, (2001) Vol. 46, No. 8, pp.
471-477. print.

ISSN: 1434-5161.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI A high-throughput SNP typing system for genome-wide association
studies.

AB. . . for genotyping. Currently available technologies require too large a
quantity of genomic DNA to genotype with hundreds or thousands of
single-nucleotide polymorphisms (SNPs

). To overcome this problem, we combined the Invader assay with
multiplex polymerase chain reaction (PCR), carried out in the
presence of antibody to Taq polymerase, as well as using a novel 384-well
card system that can reduce the required reaction volume. We amplified 100
genomic DNA fragments, each containing one SNP, in a single
tube, and analyzed each SNP with the Invader assay.

This procedure correctly genotyped 98 of the 100 SNP loci
examined in PCR-amplified samples from ten individuals; the genotypes were
confirmed by direct sequencing. The reproducibility and universality of
the method were confirmed with two additional sets of 100 SNPs.

Because we used 40 ng of genomic DNA as a template for multiplex
PCR, the amount needed to assay one SNP was only 0.4 ng;
therefore, theoretically, more than 200,000 SNPs could be
genotyped at once when 100 mug of genomic DNA is available. Our results
indicate the feasibility of undertaking. . .

L6 ANSWER 11 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

DUPLICATE 3

ACCESSION NUMBER: 2001:214630 BIOSIS

DOCUMENT NUMBER: PREV200100214630

TITLE: High-throughput multiplex SNP
genotyping with MALDI-TOF mass spectrometry: Practice,
problems and promise.
AUTHOR(S): Bray, Molly S. (1); Boerwinkle, Eric; Doris, Peter A.
CORPORATE SOURCE: (1) Human Genetics Center, University of Texas Health
Science Center at Houston, Houston, TX, 77225:
molly.s.bray@uth.tmc.edu USA
SOURCE: Human Mutation, (2001) Vol. 17, No. 4, pp. 296-304. print.
ISSN: 1059-7794.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI High-throughput multiplex SNP genotyping with
MALDI-TOF mass spectrometry: Practice, problems and promise.

AB Single nucleotide polymorphisms (SNPs) are currently being identified and mapped at a remarkable pace, providing a rich genetic resource with vast potential for disease. . . humans. High-throughput, cost effective genotyping methods are essential in order to make the most advantageous and immediate use of these SNP data. We have incorporated the use of matrix-assisted laser desorption/ionization time-of-flight mass spectrometry (MALDI-TOF) in our laboratory as a tool. . . for differentiating genotypes based on the mass of the variant DNA sequence, and have utilized this method for production scale SNP genotyping. We have combined a 4 mul PCR amplification reaction using 3 ng of genomic DNA with a secondary enzymatic. . . analysis of mini-sequencing reactions was performed using a MALDI-TOF instrument (Voyager-DE, Perseptive Biosystems, Framingham, MA). We performed both single and multiplex PCR and mini-sequencing reactions, and genotyped seven different variant sites in a random sample of 989 individuals. Genotypes generated with MS methods were compared with genotypes produced using a 5' exonuclease fluorescence-based assay (Taqman, Applied Biosystems, Foster City, CA) and a gel-based genotyping protocol. Because multiple polymorphisms can be detected in a single reaction,. . .

L6 ANSWER 12 OF 18 MEDLINE on STN DUPLICATE 4

ACCESSION NUMBER: 2001156126 MEDLINE

DOCUMENT NUMBER: 21098045 PubMed ID: 11159763

TITLE: Enabling large-scale pharmacogenetic studies by
high-throughput mutation detection and genotyping
technologies.

AUTHOR: Shi M M

CORPORATE SOURCE: Department of Applied Genomics, Genometrix Inc., The
Woodlands, TX 77381, USA.. mshi@genometrix.com

SOURCE: CLINICAL CHEMISTRY, (2001 Feb) 47 (2) 164-72. Ref: 40
Journal code: 9421549. ISSN: 0009-9147.

PUB. COUNTRY: United States
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
(REVIEW, TUTORIAL)

LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200103
ENTRY DATE: Entered STN: 20010404
Last Updated on STN: 20010404
Entered Medline: 20010322

AB . . . challenge. APPROACH: This article reviews the recent technology development in mutation detection and genotyping with a focus on genotyping of single nucleotide polymorphisms (SNPs). CONTENT: Novel mutations/polymorphisms are commonly identified by conformation-based mutation screening and direct high-throughput heterozygote sequencing. With a large amount of public sequence information available, in silico SNP mapping has also emerged as a cost-efficient way for new polymorphism identification. Gel electrophoresis-based genotyping methods for known polymorphisms include PCR coupled with restriction fragment length polymorphism analysis, multiplex PCR, oligonucleotide ligation assay, and minisequencing. Fluorescent dye-based genotyping technologies are emerging as high-throughput genotyping platforms, including oligonucleotide ligation assay, pyrosequencing, single-base extension with fluorescence detection, homogeneous solution hybridization such as TaqMan, and molecular beacon genotyping. Rolling circle amplification and Invader assays are able to genotype directly from genomic DNA without PCR amplification. DNA chip-based microarray and mass spectrometry genotyping technologies. . .

L6 ANSWER 13 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

DUPLICATE 5

ACCESSION NUMBER: 2001:125349 BIOSIS

DOCUMENT NUMBER: PREV200100125349

TITLE: An improved real time PCR method for simultaneous detection
of C282Y and H63D mutations in the HFE gene associated with
hereditary hemochromatosis.

AUTHOR(S): Walburger, D. K. (1); Afonina, I. A.; Wydro, R.

CORPORATE SOURCE: (1) Epoch Biosciences, 12277 134th Ct NE, No. 110,
Redmond,

WA, 98052: dwalburger@epochpharm.com USA

SOURCE: Mutation Research, (January, 2001) Vol. 432, No. 3-4, pp.
69-78. print.

ISSN: 0027-5107.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

AB. . . HFE gene-C282Y and H63D-are associated with greater than 90% of HH cases. We have developed a sensitive real time PCR (TaqMan) 5'-nuclease assay for single nucleotide polymorphism (SNP) detection using novel DNA chemistry, and successfully applied this method to detect these mutations. Fluorogenic PCR probes, chemically modified with a minor groove binding agent to increase duplex stability, were used in single and multiplex probe closed tube formats. The probes were tested in two commercially available thermocycling fluorimeters (the Light CyclerTM and the ABI. . .

L6 ANSWER 14 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:151558 BIOSIS

DOCUMENT NUMBER: PREV200100151558

TITLE: Enabling large-scale pharmacogenetic studies by high-throughput mutation detection and genotyping technologies.

AUTHOR(S): Shi, Michael M. (1)

CORPORATE SOURCE: (1) Genometrix Inc., 2700 Research Forest Dr., The Woodlands, TX, 77381: mshi@genometrix.com USA

SOURCE: Clinical Chemistry, (February, 2000) Vol. 47, No. 2, pp. 164-172. print.

ISSN: 0009-9147.

DOCUMENT TYPE: General Review

LANGUAGE: English

SUMMARY LANGUAGE: English

AB. . . challenge. Approach: This article reviews the recent technology development in mutation detection and genotyping with a focus on genotyping of single nucleotide polymorphisms (SNPs). Content: Novel mutations/polymorphisms are commonly identified by conformation-based mutation screening and direct high-throughput heterozygote sequencing. With a large amount of public sequence information available, in silico SNP mapping has also emerged as a cost-efficient way for new polymorphism identification. Gel electrophoresis-based genotyping methods for known polymorphisms include PCR coupled with restriction fragment length polymorphism analysis, multiplex PCR, oligonucleotide ligation assay, and minisequencing. Fluorescent dye-based genotyping technologies are emerging as high-throughput genotyping platforms, including oligonucleotide ligation assay, pyrosequencing, single-base extension with fluorescence detection, homogeneous solution hybridization such as TaqMan(R), and molecular beacon genotyping. Rolling circle amplification and InvaderTM assays are able to genotype directly from genomic DNA

without PCR amplification. DNA chip-based microarray and mass spectrometry genotyping technologies. . .

L6 ANSWER 15 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

DUPLICATE 6

ACCESSION NUMBER: 2000:360537 BIOSIS

DOCUMENT NUMBER: PREV200000360537

TITLE: Suspension arrays for high throughput, multiplexed
single nucleotide polymorphism
genotyping.

AUTHOR(S): Armstrong, Barbara; Stewart, Michael; Mazumder, Abhijit (1)

CORPORATE SOURCE: (1) Motorola Biochip Systems, 4088 Commercial Ave.,
Northbrook, IL, 60062 USA

SOURCE: Cytometry, (June 1, 2000) Vol. 40, No. 2, pp. 102-108.
print.

ISSN: 0196-4763.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI Suspension arrays for high throughput, multiplexed
single nucleotide polymorphism genotyping.

AB. . . Background: Genetic diversity can help explain disease
susceptibility and differential drug response. The most common type of
variant is the single nucleotide polymorphism
(SNP). We present a low-cost, high throughput assay for
SNP genotyping. Methods: The assay uses oligonucleotide probes
covalently attached to fluorescently encoded microspheres. These probes
are hybridized directly to fluorescently. . . in a standard flow
cytometer. Results: The genotypes determined with our assay are in good
agreement with those determined by TaqMan. The range of G/C
content for oligonucleotide probes was 23.5-65% in the 17 bases
surrounding the SNP. Further optimization of probe length and
target concentration is shown to dramatically enhance the assay
performance for certain SNPs. Using microspheres which have
unique fluorescent signatures, we performed a 32-plex assay where we
simultaneously determined the genotypes of eight different polymorphic
genes. Conclusions: We demonstrate, for the first time, the feasibility of
multiplexed genotyping with suspension arrays using direct
hybridization analyses. Our approach enables probes to be removed from or
added to an array, enhancing flexibility over conventional chips. The
ability to multiplex both the PCR preparation and the
hybridization should enhance the throughput, cost, and speed of the assay.

L6 ANSWER 16 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

ACCESSION NUMBER: 2001:528511 BIOSIS

DOCUMENT NUMBER: PREV200100528511

TITLE: A single-tube primer extension reaction enabling
multiplex SNP interrogation.

AUTHOR(S): Wei, Dong (1); Johnson, Martin (1); Kuo, Sophia (1);
Hanachi, Parisa (1); Yu, Wendy (1); Dong, Penny (1)

CORPORATE SOURCE: (1) Applied Biosystems, Foster City, CA USA

SOURCE: International Genome Sequencing and Analysis Conference,
(2000) Vol. 12, pp. 96-97. print.

Meeting Info.: 12th International Genome Sequencing and
Analysis Conference Miami Beach, Florida, USA September
12-15, 2000

DOCUMENT TYPE: Conference

LANGUAGE: English

SUMMARY LANGUAGE: English

TI A single-tube primer extension reaction enabling multiplex
SNP interrogation.

AB A variety of methods may be used to characterize and screen for
single nucleotide polymorphisms. Analysis
platforms include microarray scanning, real time PCR analysis (Taqman) and MALDI-TOF. Additionally, electrophoresis based techniques include OLA analysis, dideoxy sequencing and single-nucleotide primer extension (e.g., SNaPshot). SNaPshot is a single-tube reaction designed for elucidation of individual loci within known sequence contexts for the purpose of SNP screening. The reaction is designed as a premix that contains all of the components except primers and templates. The completed. . . identifies one nucleotide located 3' relative to the primer site. We have reformulated our SNaPshot reagent mix to enable robust multiplex SNP interrogation against multiple templates in varying amounts. The resulting multiple products can then be analyzed by electrophoresis in the presence. . .

L6 ANSWER 17 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

ACCESSION NUMBER: 2001:494471 BIOSIS

DOCUMENT NUMBER: PREV200100494471

TITLE: Multiplexed genotyping assay that combines a
novel labeling strategy with microchannel electrophoresis.

AUTHOR(S): Cronin, Maureen T. (1); Williams, Stephen J. (1); Pho,
Mylan (1); Wei, Jing (1); Leon, Suzan (1); Matray, Tracy
(1); Singh, Sharat (1); Livak, Ken J.; Dong, Penny;
Mansfield, Elaine S. (1)

CORPORATE SOURCE: (1) ACLARA BioSciences Inc., Mountain View, CA USA

SOURCE: International Genome Sequencing and Analysis Conference,
(2000) Vol. 12, pp. 27. print.

Meeting Info.: 12th International Genome Sequencing and

Analysis Conference Miami Beach, Florida, USA September
12-15, 2000

DOCUMENT TYPE: Conference

LANGUAGE: English

SUMMARY LANGUAGE: English

TI Multiplexed genotyping assay that combines a novel labeling
strategy with microchannel electrophoresis.

AB Genome-wide SNP scans to support large association studies and
specific genotyping panels used for phenotype prediction are both
requirements of new technologies. . . to comprehensively address
emerging genotyping demands. By combining ACLARA's proprietary probe
labeling strategy with existing, robust genotyping biochemistries such as
TaqMan>TM we have designed a flexible, multiplexed assay
format for analyzing nucleic acids in a broad spectrum of genomics needs.
Genotyping applications using this assay format are. . . This assay
configuration is uniquely characterized by its modular composition. Sets
of electrophoretic mobility tags (e-TagsTM) "code" specific probes in
multiplexed sets. These tags are released during amplification via
TaqMan cleavage and subsequently separated by capillary
electrophoresis. The electrophoretic pattern is "decoded" to yield a
genotype. Initially, 20 e-Tags will be used to provide 10-plex genotyping
capability during each capillary separation. The modular assay design
allows a single genotyping multiplex to be applied to a large
number of samples or, alternatively, parallel multiplex
reactions can be done to assay many polymorphisms in a single sample.
Multiplexed e-Tag genotyping data for the pharmacogenetics targets
CYP2D6 and ApoE will be presented.

L6 ANSWER 18 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS
INC. on STN

DUPLICATE 7

ACCESSION NUMBER: 1999:369986 BIOSIS

DOCUMENT NUMBER: PREV199900369986

TITLE: High-throughput genotyping method for glutathione
S-transferase T1 and M1 gene deletions using TaqMan
(R) probes.

AUTHOR(S): Shi, Michael M. (1); Myrand, Scott P.; Bleavins, Michael
R.; de la Iglesia, Felix A.

CORPORATE SOURCE: (1) Pathology and Experimental Toxicology, Parke-Davis
Pharmaceutical Research, Warner-Lambert Company, 2800
Plymouth Rd., Ann Arbor, MI, 48105 USA

SOURCE: Research Communications in Molecular Pathology and
Pharmacology, (Jan., 1999) Vol. 103, No. 1, pp. 3-15.
ISSN: 1078-0297.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI High-throughput genotyping method for glutathione S-transferase T1 and M1 gene deletions using TaqMan(R) probes.

AB. . . glutathione-S-transferase theta and mu (GSTT1 and GSTM1). This method utilizes the 5'-nuclease activity of Taq polymerase in conjunction with fluorogenic TaqMan(R) probes. In contrast to traditional allelic discrimination genotyping to detect single nucleotide polymorphisms, the current assay has been designed to detect gene deletion by utilizing custom-designed TaqMan probes in conjunction with an exogenous internal positive control probe. The TaqMan genotyping results were validated by a commonly used multiplex PCR technique. Screening of 71 unrelated individuals revealed gene deletion (null) genotype of 15.5% and 40.8% for GSTT1 and GSTM1, respectively. This TaqMan genotyping method is rapid, reproducible, and highly sensitive and could be applied toward fully automated large-scale genotyping.